In the Claims

Please amend the claims presented during the international phase as follows.

Applicant presents a full set of claims showing markups of the claims with insertions and deletions indicated by underlining (or double bracketing) and strikethrough text, respectively.

- 1. (Original) A method for delivering an anti-target compound to a subject for the treatment of a disorder without being inactivated by administering thereto a polypeptide construct comprising one or more single domain antibodies directed against said target.
- 2. (Currently amended) A method according to claim 1 wherein said target is located in the gut system, and said [[a]] polypeptide construct is delivered orally.
- 3. (Currently amended) A method according to claim 1 wherein said target is located in vaginal and/or rectal tract, and said [[a]] polypeptide construct is delivered to the vaginal and/or rectal tract.
- 4. (Currently amended) A method according to claim 1 wherein said target is located in nose, upper respiratory tract and/or lung, and said [[a]] polypeptide construct is delivered to nose, upper respiratory tract and/or lung.
- 5. (Currently amended) A method according to claim 1 wherein said target is located in intestinal mucosa, and said [[a]] polypeptide construct is delivered orally.
- 6. (Currently amended) A method according to claim 1 wherein said target is located in the tissues beneath the tongue, and said [[a]] polypeptide construct is delivered to the tissues beneath the tongue.
- 7. (Currently amended) A method according to claim 1 wherein said target is located in the skin, and said [[a]] polypeptide construct is delivered topically.
- 8. (Currently amended) A method according to claim 1 wherein said target is in, or accessible via the blood, and said [[a]] polypeptide construct is delivered orally, to the

vaginal and/or rectal tract, nasally, by inhalation though the mouth or nose, to the tissues beneath the tongue, or topically.

4

9. A polypeptide construct comprising at least one single domain antibody directed against a target, wherein the single domain antibody corresponds to a sequence represented by any of SEQ ID NOs: 12 to 70, for use in treating, preventing and/or alleviating the symptoms of disorders which are susceptible to modulation by an anti-target therapeutic compound that is able pass through the gastric environment without being inactivated.

10.-14. (Canceled)

- 15. (Currently amended) A method according to <u>claim 1</u> any of claims 1 to 8 or polypeptide construct according to any of claim 9 to 14, wherein said target is TNF-alpha and the disorder is inflammation.
- 16. (Currently amended) A method or polypeptide according to claim 15, wherein [[a]] the single domain antibody corresponds to a sequence represented by any of SEQ ID NOs: 12 to 14.
- 17. (Currently amended) A method according to <u>claim 1</u> any of claims 1 to 8 or polypeptide construct according to any of claim 9 to 14, wherein said target is CEA and the disorder <u>is</u> colon cancer.
- 18. (Currently amended) A method according to <u>claim 1</u> any of claims 1 to 8 or polypeptide construct according to any of claim 9 to 14, wherein said target is EGFR and the disorder is any of head, neck, lung and colon cancer.
- 19. (Currently amended) A method or polypeptide construct according to claim 18, wherein [[a]] the single domain antibody corresponds to a sequence represented by any of SEQ ID NOs: 23 to 44.
- 20. (Currently amended) A method according to <u>claim 1</u> any of claims 1 to 8 or polypeptide construct according to any of claim 9 to 14, wherein said target is antigen of *Helicobacter pylori* and the disorder is any of indigestion[[,]] and gastritis.

- 5
- 21. (Currently amended) A method according to <u>claim 1</u> any of claims 1 to 8 or polypeptide construct according to any of claim 9 to 14, wherein said target is antigen of *Mycobacterium tuberculosis* and the disorder is tuberculosis.
- 22. (Currently amended) A method according to <u>claim 1</u> any of claims 1 to 8 or polypeptide construct according to any of claim 9 to 14, wherein said target is antigen of *influenza* virus and the disorder is flu.
- 23. (Currently amended) A method according to <u>claim 1</u> any of claims 1 to 8 or polypeptide construct according to any of claim 9 to 14, wherein said target is antigen of MMP and the disorder is cancer.
- 24. (Currently amended) A method or polypeptide construct according to claim 23, wherein [[a]] the single domain antibody corresponds to a sequence represented by any of SEQ ID NOs: 15 to 22.
- 25. (Currently amended) A method according to <u>claim 1</u> any of claims 1 to 8 or polypeptide construct according to any of claim 9 to 14, wherein said target is <u>an</u> antigen of IFN-gamma and the disorder is any of cancer, transplant rejection, auto immune disorder.
- 26. (Currently amended) A method or polypeptide construct according to claim 25, wherein [[a]] the single domain antibody corresponds to a sequence represented by any of SEQ ID NOs: 45 to 70.
- 27. (Currently amended) A method according to <u>claim 1</u>, <u>any of claims 1 to 8 or</u> polypeptide construct according to any of claim 9 to 14 wherein said target is any of <u>an</u> antigen of *Helicobacter pylori*, <u>an</u> antigen of *Mycobacterium tuberculosis*, <u>and an</u> antigen of *influenza* virus.
- 28. (Currently amended) A polypeptide construct comprising (a) at least one single domain antibody directed against an internalising cellular receptor, and (b) at least one single domain antibody directed against a therapeutic target, or at least one therapeutic polypeptide or agent.

- 29. (Canceled)
- 30. (Currently amended) A polypeptide construct according to <u>claim 28 elaims 28 or 29</u> wherein said internalising cellular receptor is Epidermal Growth Factor receptor.
- 31. (Original) A polypeptide construct according to claim 30 wherein a single domain antibody directed against an internalising cellular receptor corresponds to a sequence represented by SEQ ID NO: 23 to 44.
- 32. (Currently amended) A polypeptide construct according to <u>claim 28 elaims 28 or 29</u> wherein said internalising cellular receptor is any of LDL receptor, FGF2[[r]] receptor, ErbB2[[r]] receptor, transferring transferring receptor, <u>PDGr PDGF receptor</u>, <u>VEGr VEGF receptor</u>, or PsmAr.
- 33. (Currently amended) A polypeptide construct according to <u>claim 28</u> any of claims 28 to 32 wherein a single domain antibody directed against a therapeutic target, is directed against PDK1.
- 34. (Canceled)
- 35. (Currently amended) A polypeptide construct according to <u>claim 28</u> any of claims 28 to 32 wherein a single domain antibody directed against a therapeutic target is directed against any of GSK1, Bad, caspase and Forkhead.
- 36. (Canceled)
- 37. (Original) Method for delivering an anti-target therapeutic compound to the interior of a cell comprising administering to a subject a polypeptide construct according to claim 28 any of claims 28 to 36.
- 38. (Canceled)

Int'l Application No.: PCT/BE03/00190 7 Int'l Filing Date: 7 November 2003

39. (Currently amended) A method according to claim <u>37</u> [[38]] wherein said cell is located in the gut system, and said [[a]] polypeptide construct is delivered orally.

- 40. (Currently amended) A method according to claim <u>37</u> [[38]] wherein said cell is located in vaginal and/or rectal tract, and said [[a]] polypeptide construct is delivered to the vaginal and/or rectal tract.
- 41. (Currently amended) A method according to claim <u>37</u> [[38]] wherein said cell is located in nose, upper respiratory tract and/or lung, and said [[a]] polypeptide construct is delivered to nose, upper respiratory tract and/or lung.
- 42. (Currently amended) A method according to claim <u>37</u> [[38]] wherein said cell is located in intestinal mucosa, and said [[a]] polypeptide construct is delivered orally.
- 43. (Currently amended) A method according to claim <u>37</u> [[38]] wherein said cell is located in the tissues beneath the tongue, and said [[a]] polypeptide construct is delivered to the tissues beneath the tongue.
- 44. (Currently amended) A method according to claim <u>37</u> [[38]] wherein said cell is located in the skin, and said [[a]] polypeptide construct is delivered topically.
- 45. (Currently amended) A method according to claim <u>37</u> [[38]] wherein said cell is in, or accessible via the blood, and said [[a]] polypeptide construct is delivered orally, to the vaginal and/or rectal tract, nasally, by inhalation though the mouth or nose, to the tissues beneath the tongue, or topically.
- 46. (Currently amended) A polypeptide construct according to any of claims 9 or 28 to 14, 28 to 36, or as used in a method according to claim 1 any of claims 1 to 8, 15 to 27, 37 to 45, wherein the single domain antibodies are humanized Camelidae VHHs, an homologous sequence, a functional portion, or a functional portion of an homologous sequence of the full length single domain antibody, or wherein the polypeptide construct is an homologous sequence, a functional portion, or a functional portion of an homologous sequence of the full length polypeptide construct.

Int'l Application No.: PCT/BE03/00190 8 Int'l Filing Date: 7 November 2003

47.-49. (Canceled)

50. (Currently amended) A nucleic acid encoding a polypeptide construct according to any of claims 9 to 14, 28 to 36, or 46 to 49.

- 51. (Currently amended) A composition comprising a polypeptide construct according to any of claims 9, 28 or 46 as defined in any of the preceding claims, together with a pharmaceutical carrier.
- 52. (New) A method according to claim 1, wherein said target is antigen of IgE and the disorder is allergic response.
- 53. (New) A method according to claim 52, wherein a single domain antibody corresponds to a sequence represented by any of SEQ ID NOs: 1 to 11.
- 54. (New) A polypeptide construct comprising at least one single domain antibody directed against IgE.
- 55. (New) A polypeptide construct according to claim 54 wherein at least one single domain antibody is a *Camelidae* VHH.
- 56. (New) A polypeptide construct according to claim 54 wherein at least one single domain antibody corresponds to a sequence represented by any of SEQ ID NOs: 1 to 11.
- 57. (New) A polypeptide construct according to claim 54, wherein the number of anti-IgE single domain antibodies is at least two.
- 58. (New) A polypeptide construct according to claim 57, wherein at least one single domain antibody is a humanized *Camelidae* VHH.
- 59. (New) A polypeptide construct according to claim 54, wherein a single domain antibody is an homologous sequence, a functional portion, or a functional portion of an homologous sequence of the full length single domain antibody.

Int'l Application No.: PCT/BE03/00190 9 Int'l Filing Date: 7 November 2003

60. (New) A polypeptide construct according to claim 54, wherein the polypeptide construct is an homologous sequence, a functional portion, or a functional portion of an homologous sequence of the full length polypeptide construct.

- 61. (New) A nucleic acid encoding a polypeptide construct according to claim 54.
- 62. (New) A method for treating and/or preventing and/or alleviating disorders relating to inflammatory processes comprising administering to a subject in need of such treatment a polypeptide construct according to claim 54.
- 63. (New) A composition comprising a polypeptide construct according to claim 54 together with a pharmaceutical carrier.